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Wu et al.

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(54) **AC GATE ION FILTER METHOD AND APPARATUS**

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(60) Provisional application No. 61/784,324, filed on Mar. 14, 2013, provisional application No. 61/801,722, filed on Mar. 15, 2013.

(51) **Int. Cl.**
H01J 49/40 (2006.01)
H01J 49/06 (2006.01)
C07B 63/00 (2006.01)
G01N 27/62 (2006.01)

(52) **U.S. Cl.**

CPC **H01J 49/061** (2013.01); **C07B 63/00** (2013.01); **G01N 27/622** (2013.01)

(58) **Field of Classification Search**

USPC 250/281, 282, 283, 286, 287
See application file for complete search history.

(56) **References Cited**

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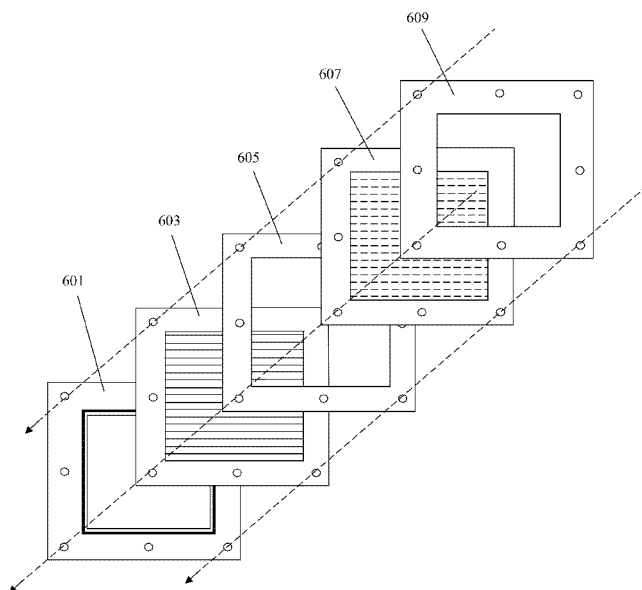
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Primary Examiner — Nicole Ippolito

(57) **ABSTRACT**

The present invention uses an AC voltage instead of DC voltage on an ion gate to filter/selectively pass ions. The ions that pass through the AC ion gate can be further separated in a spectrometric instrument. An ion mobility spectrometer using the AC ion gate can achieve better gating performance. For a time of flight ion mobility spectrometer with an AC ion gate, a narrow pulse of selected ions can be passed into a drift tube where they are separated based on their low field ion mobility. Moreover, when the AC voltage at the AC ion gate has a waveform as used for differential ion mobility spectrometry, the time of flight ion mobility spectrometer is converted into a two dimensional separation spectrometer, where ions are first separated based on their high field ion mobility and then further separated based on their low field ion mobility.

16 Claims, 13 Drawing Sheets



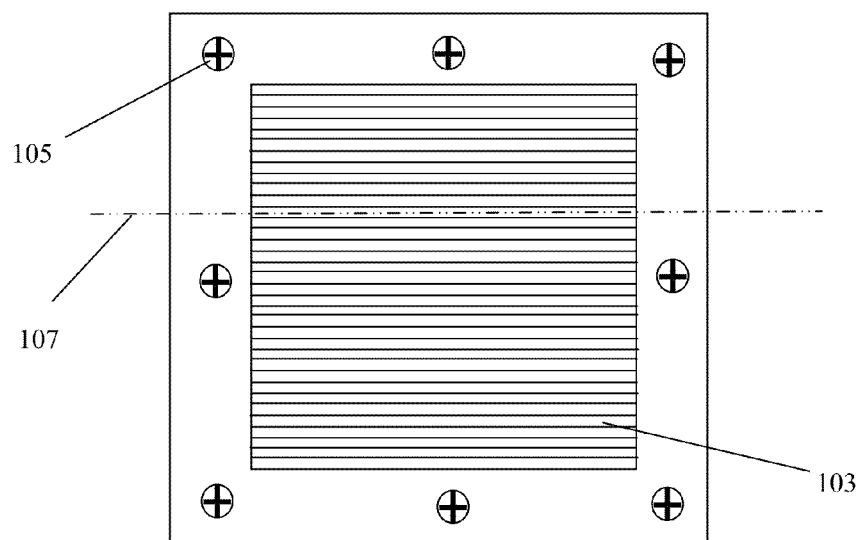


Figure 1

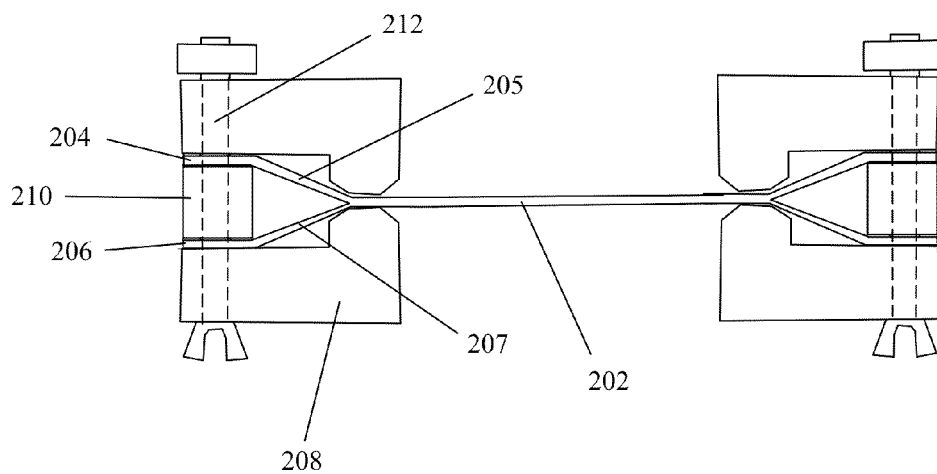


Figure 2

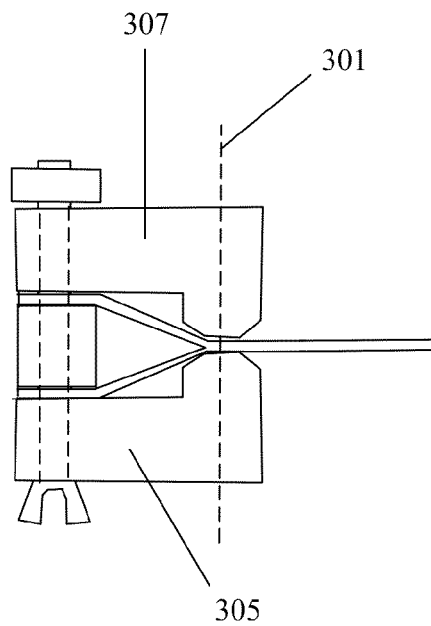


Figure 3A

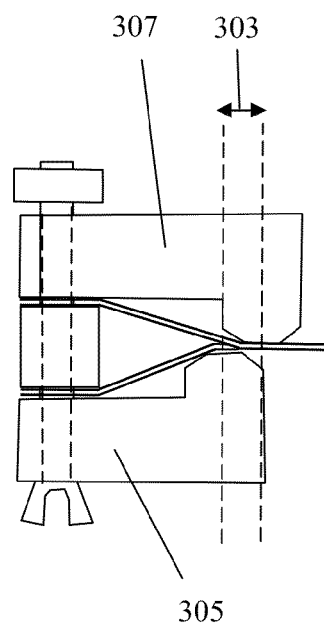


Figure 3B

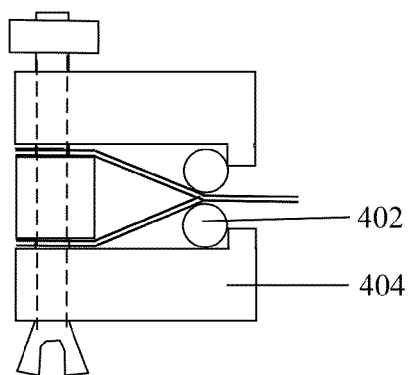


Figure 4

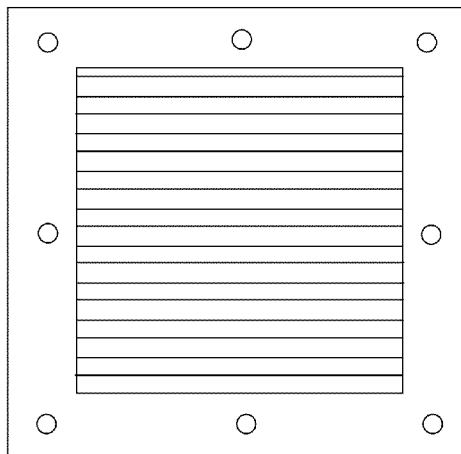


Figure 5A

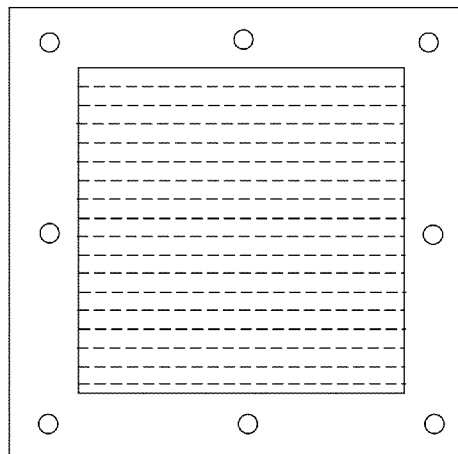


Figure 5B

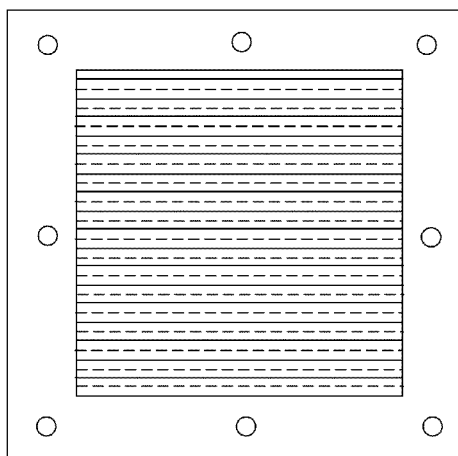


Figure 5C

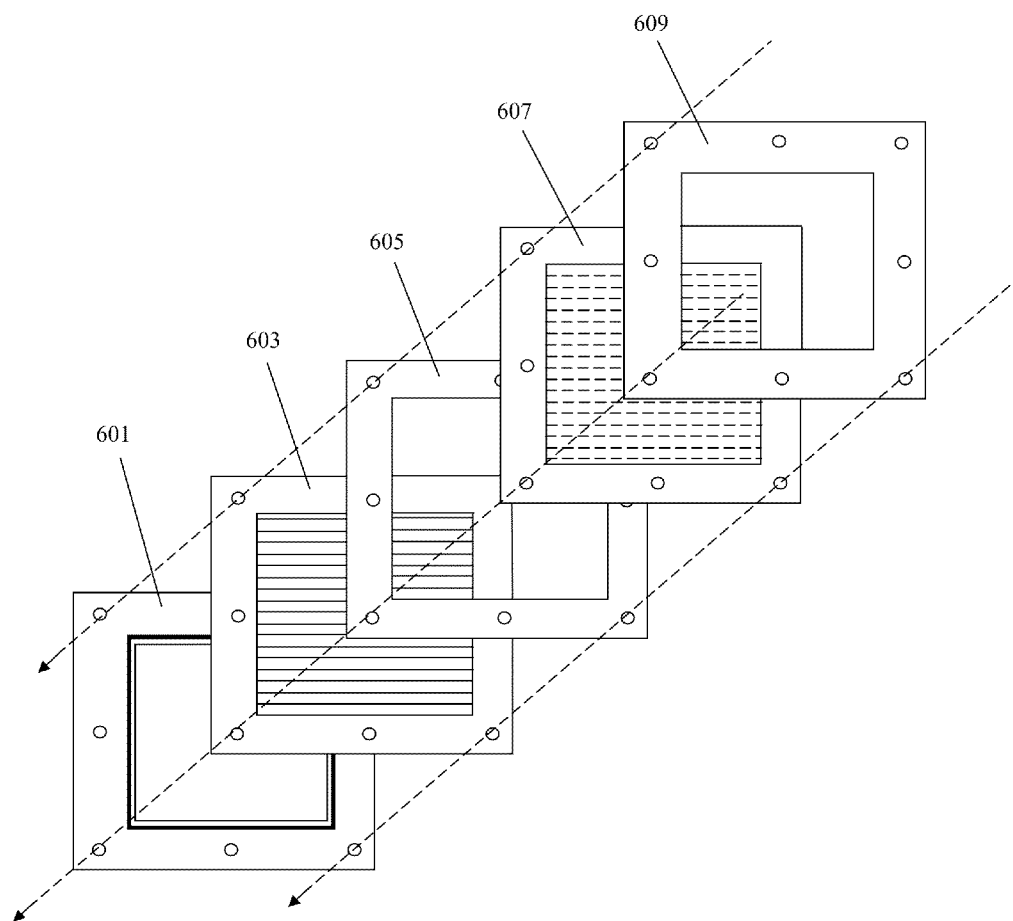


Figure 6

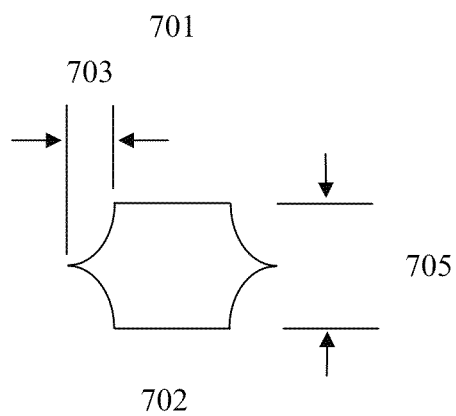


Figure 7A

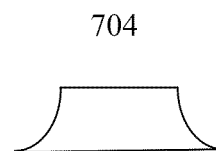


Figure 7B

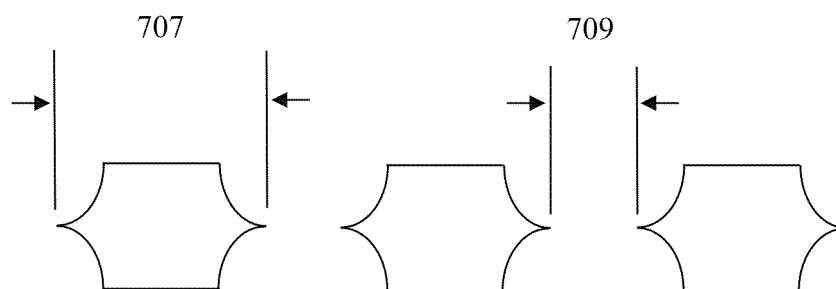


Figure 7C

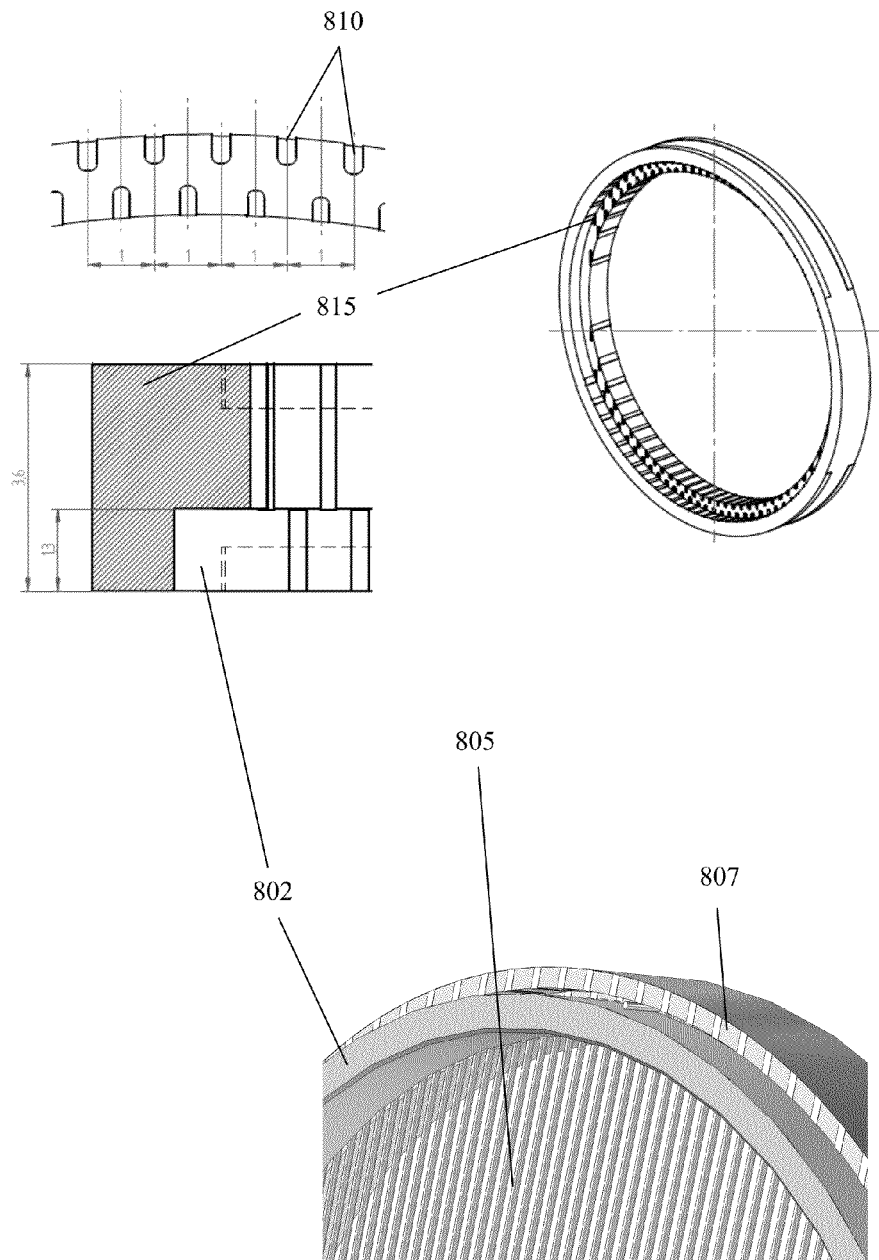


Figure 8

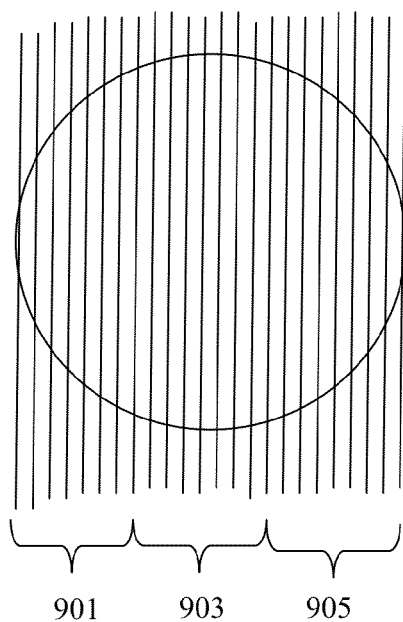


Figure 9

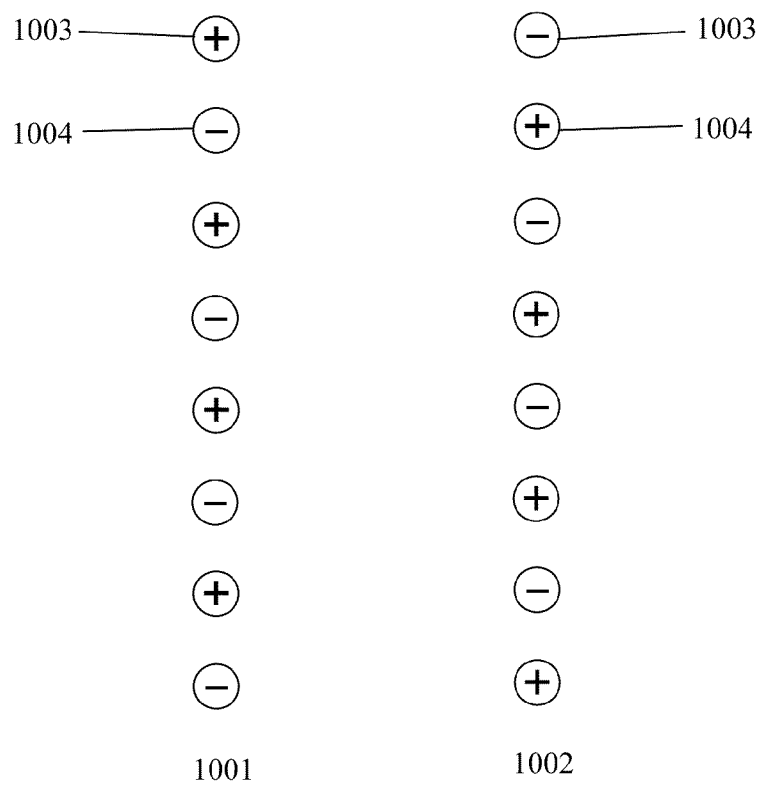


Figure 10

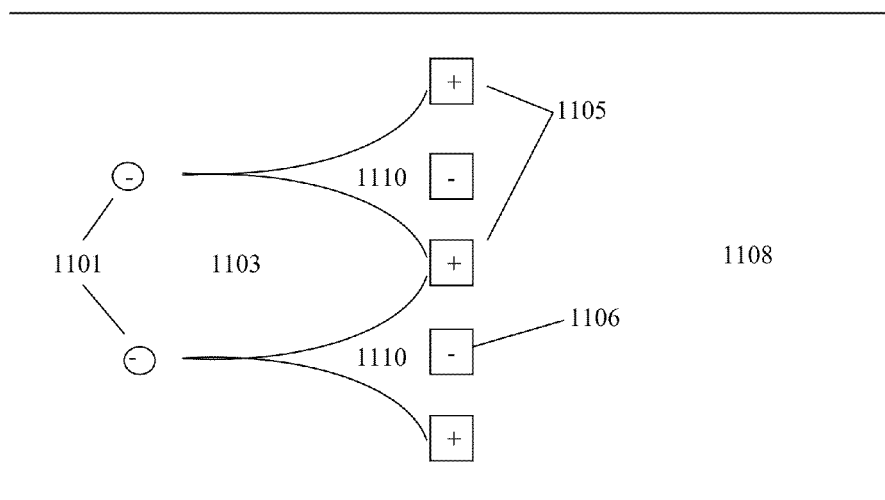


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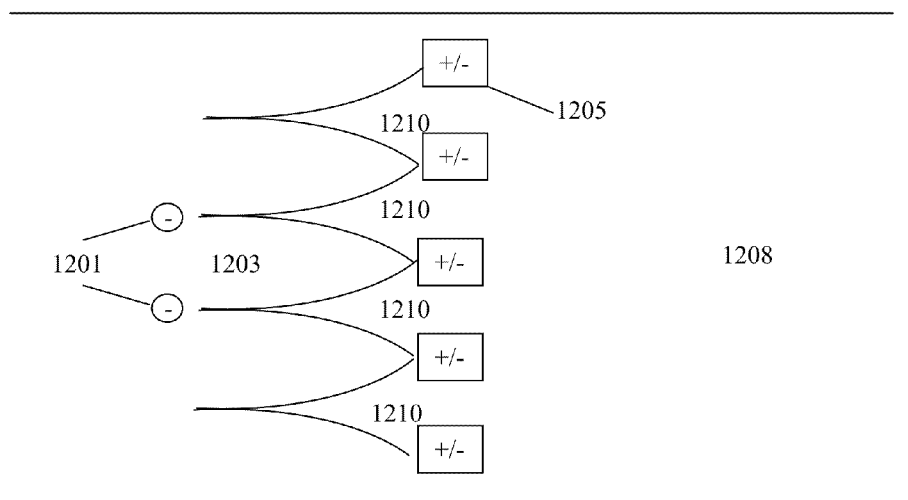


Figure 12

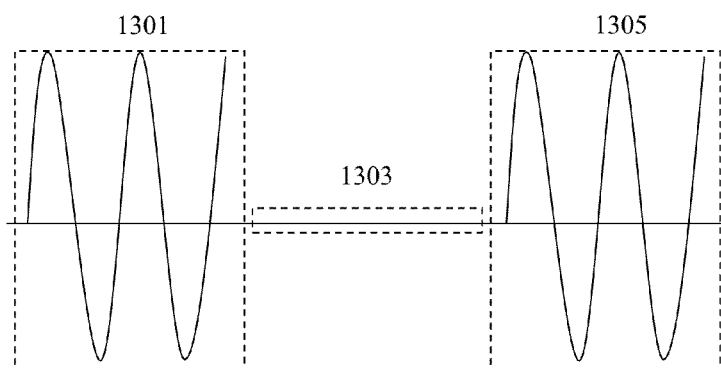


Figure 13

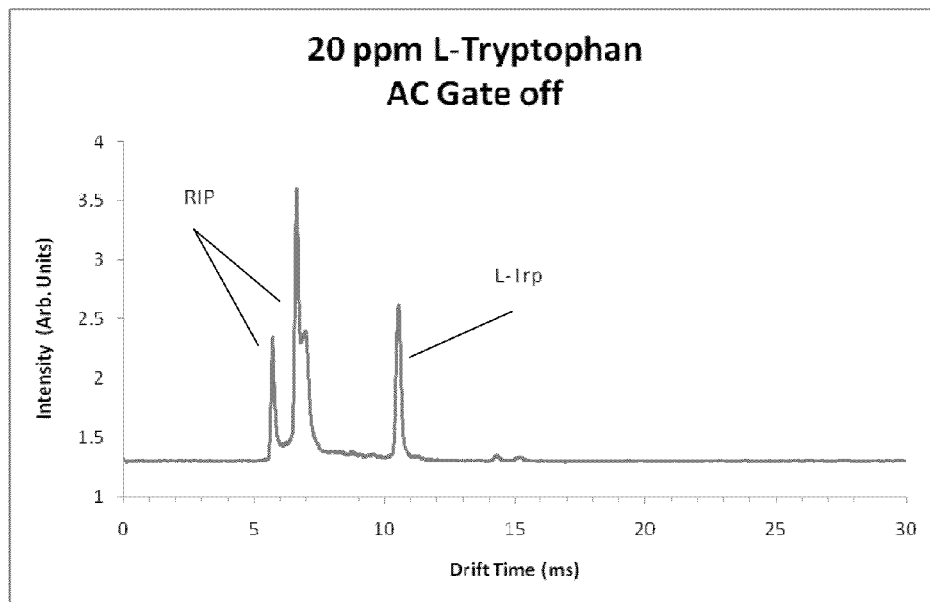


Figure 14

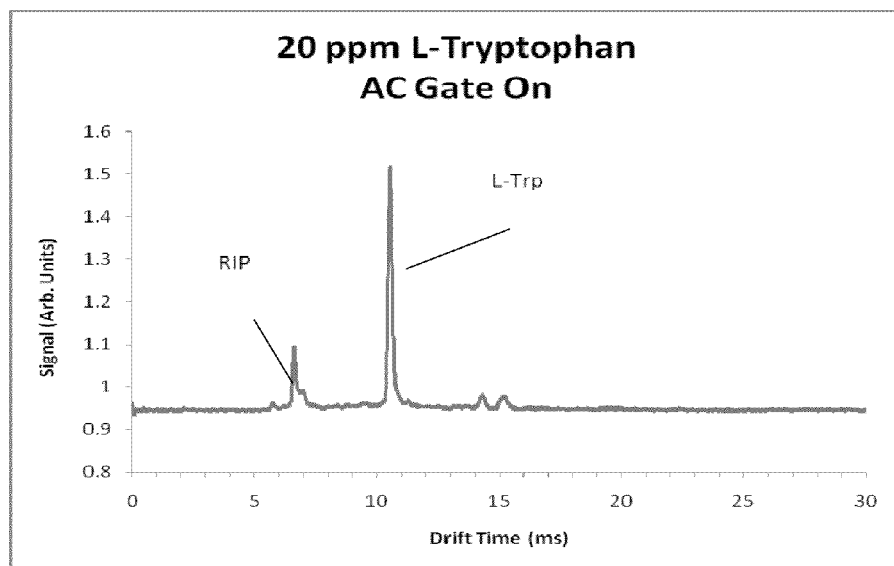


Figure 15

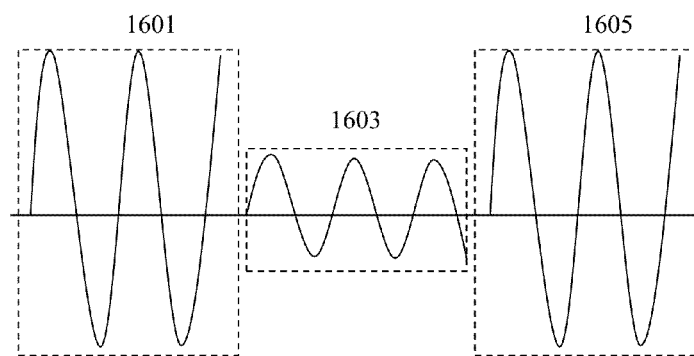


Figure 16

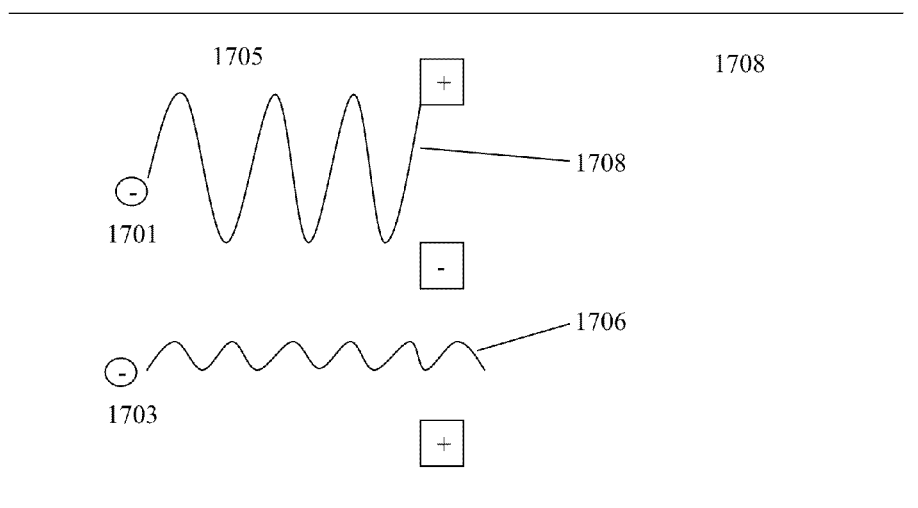


Figure 17

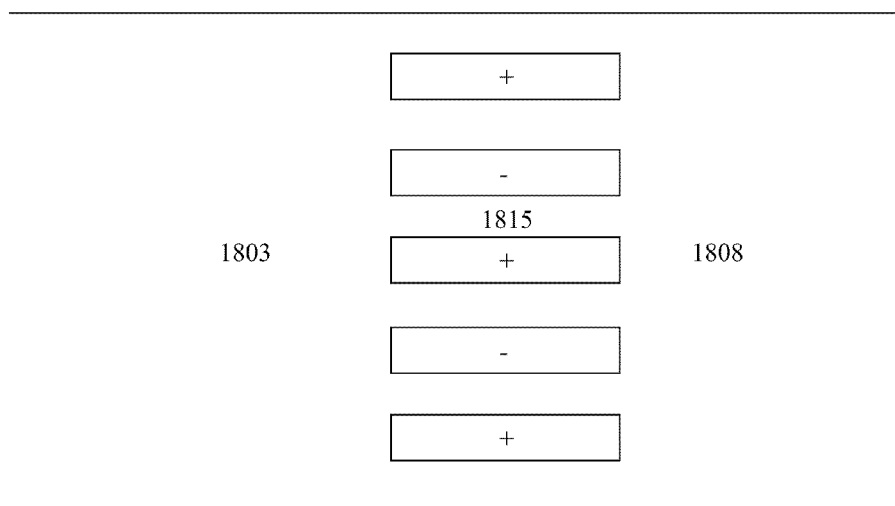


Figure 18

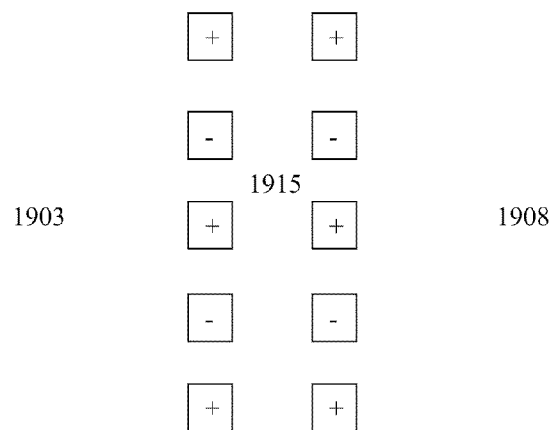


Figure 19

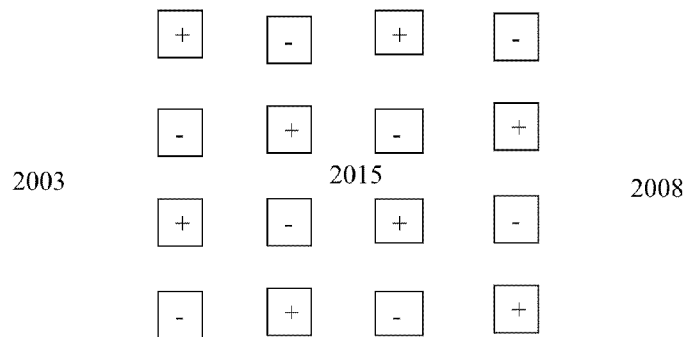


Figure 20

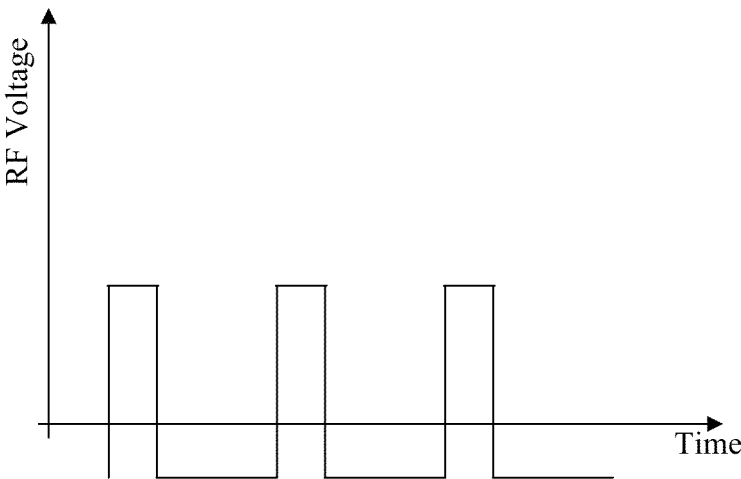


Figure 21

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AC GATE ION FILTER METHOD AND APPARATUS

The present application claims the benefit of and priority to corresponding U.S. Provisional Patent Application No. 61/784,324, filed on Mar. 14, 2013 and 61/801,722, filed on Mar. 15, 2013 and continuation in part of U.S. patent application Ser. No. 12/763,092, respectively, the entire content of the applications are herein incorporated by reference.

BACKGROUND OF THE INVENTION

Many analytical instruments, such as ion mobility spectrometers (IMS), can require a gating device for turning on and off a flowing stream of ions and/or other charged particles. IMS are widely used in field chemical analysis. IMS separate ionic species based on their ion mobility in a given media (either gas or liquid). Recent development of the IMS technology results in two forms of IMS instruments and systems. The time-of-flight (TOF) IMS separate ions based on their steady state ion mobilities under constant electric field. High resolving power with IMS has been achieved with the TOF-IMS instruments. Alternatively, devices that separate ions based their mobility changes under high field conditions, such as field asymmetric ion mobility spectrometer (FAIMS) or differential mobility spectrometer (DMS), can also be used.

Even though the gating device is a minor component in the overall design of an IMS, if manufactured correctly, this component can improve the IMS resolution and system performance. The gating device is used to regulate the injection of ion packets into the analytical instrument. There are many deficiencies with the current approaches for manufacturing gating devices.

Traditionally the gating device has been used to regulate the injection of ion packets into the analytical instrument. Even though the gating device is a minor component in the overall design of an IMS, this device is an important part that can improve the level resolution between peaks in the IMS by providing a compact ion packet without significant diffusion. Many inventions have been proposed around the manufacturing and designing the gating device for improving the resolution without major improvements. The present invention modifies the gating device in a manner that improves peak resolution and is able to control which size ions are injected into the analytical instrument. This novel gating device significantly reduces the analysis of complex samples with multiple components such that lower mobility ions are not able to enter the drift tube.

SUMMARY OF THE INVENTION

The present invention generally relates to systems and methods for transmitting beams of charged particles, and in particular to such systems and methods that employ deflecting at least one set of grid elements into the same plane, such that the grid elements are interleaved.

In one embodiment of the present invention, at least one electrically substrate (conducting or non-conducting) is used to deflect at least one set of the grid elements into the same plane, such that the grid elements are interleaved. The ion gate has a first and second set of electrically isolated grid elements that lie in the same plane where the respective sets of grid elements are applied to alternate potentials. The advanced grid manufacturing methods and features are disclosed.

The present invention generally relates to systems and methods for transmitting beams of charged particles, and in

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particular to such systems and methods that let only a portion of ions to the drift tube of the IMS by employing an AC voltage on the gate wires. By using an AC voltage there is a reduction in the size of the ion depletion area in front of the gate when it is closed, thereby providing a higher peak resolution.

BRIEF DESCRIPTION OF THE DRAWINGS

The foregoing and other aspects, embodiments, and features of the inventions can be more fully understood from the following description in conjunction with the accompanying drawings. In the drawings like reference characters generally refer to like features and structural elements throughout the various figures. The drawings are not necessarily to scale, emphasis instead being placed upon illustrating the principles of the inventions.

FIG. 1 shows the completed ion gate from a front view.

FIG. 2 shows a cross-sectional top view of the ion gate.

FIGS. 3A and 3B show two different methods to deflect the grid elements.

FIG. 4 shows an alternative method to deflect the grid elements.

FIGS. 5A-5C illustrates the respective sets of gate elements.

FIG. 6 illustrates the process of manufacturing the ion gate.

FIG. 7A-7C shows the photo etched edges of the grid elements.

FIG. 8 schematically shows a construction of a Bradbury-Nielsen ion gate using metalized dielectric rings and parallel wires.

FIG. 9 is a schematic example of a segmented Bradbury-Neilson gate.

FIG. 10 illustrates an operation method of an ion gate.

FIG. 11 shows an example of a gate using DC voltage.

FIG. 12 shows an example of a gate using AC voltage.

FIG. 13 shows the AC gate in the closed position and then completely open and then closed again.

FIG. 14 shows an example of an IMS spectrum using 20 parts per million (ppm) of an L-Tryptophan sample in 80/20 methanol and water using a DC voltage on the ion gate.

FIG. 15 shows an example of an IMS spectrum using 20 parts per million (ppm) of an L-Tryptophan sample in 80/20 methanol and water using a AC voltage on the ion gate.

FIG. 16 shows the AC mobility selecting gate in the closed position having a voltage of 240 and then partially open having a voltage of 100 and then closed again having a voltage of 240.

FIG. 17 shows the trajectory of ions using a 67 kHz AC frequency.

FIG. 18 shows gate wires that have a large surface area.

FIG. 19 shows 2 sets of gate wires aligned in phase.

FIG. 20 shows 4 sets of gate wires in an array that are out of phase.

FIG. 21 shows a typical asymmetric waveform used for differential ion mobility spectrometer or field asymmetric ion mobility spectrometer

DETAILED DESCRIPTION OF VARIOUS EMBODIMENTS

The term ion mobility separator, and ion mobility spectrometer, and ion mobility based spectrometers are used interchangeably in this invention, often referred to as IMS, including time-of-flight (TOF) IMS, differential mobility spectrometers (DMS), field asymmetric ion mobility spectrometers (FAIMS) and their derived forms. A time of flight

ion mobility spectrometer and their derived forms refers to, in its broadest sense, any ion mobility based separation device that characterize ions based on their time of flight over a defined distance. A FAIMS, a DMS, and their derived forms separate ions based on their ion mobility characteristics under high values of normalized electric field.

The systems and methods of the present inventions may make use of "drift tubes." The term "drift tube" is used herein in accordance with the accepted meaning of that term in the field of ion mobility spectrometry. A drift tube is a structure containing a neutral gas through which ions are moved under the influence of an electrical field. It is to be understood that a "drift tube" does not need to be in the form of a tube or cylinder. As understood in the art, a "drift tube" is not limited to the circular or elliptical cross-sections found in a cylinder, but can have any cross-sectional shape including, but not limited to, square, rectangular, circular, elliptical, semi-circular, triangular, etc. In many cases, a drift tube is also referred to the ion transportation and/or ion filter section of a FAIMS or DMS device.

Neutral gas is often referred to as a carrier gas, drift gas, buffer gas, etc. and these terms are considered interchangeable herein. The gas is at a pressure such that the mean free path of the ion, or ions, of interest is less than the dimensions of the drift tube. That is the gas pressure is chosen for viscous flow. Under conditions of viscous flow of a gas in a channel, conditions are such that the mean free path is very small compared with the transverse dimensions of the channel. At these pressures the flow characteristics are determined mainly by collisions between the gas molecules, i.e. the viscosity of the gas. The flow may be laminar or turbulent. It is preferred that the pressure in the drift tube is high enough that ions will travel a negligible distance, relative to the longitudinal length of the drift tube, therefore a steady-state ion mobility is achieved.

The phrase "and/or," as used herein in the specification and in the claims, should be understood to mean "either or both" of the elements so conjoined, i.e., elements that are conjunctively present in some cases and disjunctively present in other cases.

Unless otherwise specified in this document the term "particle" is intended to mean chemical and/or biological single or plurality of sub-atomic particle, atom, molecule, large or macro molecule, nanoparticle, or other matters that are vapor, droplets, aerosol, liquid, solid that follow a mobile medium, where the medium can be a gas, a liquid, supercritical fluid and/or other fluidic materials.

The present invention generally relates to systems and methods for transmitting beams of charged particles, and in particular to such systems and methods that employ deflecting at least one set of grid elements into the same plane.

As used herein, the term "grid element" generally refers to wire, rod, cable, thin metal foil piece that can be planar, square, rectangular, circular, elliptical, semi-circular, triangular, but not limited to these examples. The grid element can be made of any electrically conducting material.

The term "gate element" generally refers to a structure that includes one or more grid elements that can be spatially arranged with a gap between each other.

One aspect of the invention relates to manufacturing an ion gate in such a way that the ion gate can be produced in a simple, reproducible, and reliable manner. FIG. 6 illustrates a non-limiting process for manufacturing the ion gate. This method for manufacturing an ion gate for a charged particle stream begins with the fabrication of the gate elements **603** and **607** that includes the grid elements. Each of the gate elements **603** and **607** are made so that the two different pairs

of grid elements can be interleaved without contacting each other. Followed by assembling an insulating layer (electrically non-conductive, the insulating layer can be any size and shape, such as a square or a disk with openings, or a washer, that allows a different potential to the gate elements) **605** between the gate elements **603** and **607** to electrically isolate the gate elements. The substrate throughout this patent is typically non-conducting, but can also be a conducting material for some applications. Then two electrically non-conductive substrates **601** and **609** are added to deflect the grid elements in the gate elements into the same plane. FIG. 6 shows a non-limiting example, wherein two electrically non-conductive substrates are used. Similarly, one electrically non-conductive substrate can also be used to manufacture the ion gate. Finally, the gate elements are secured together along with the electrically non-conductive substrates and the insulating layer. In an alternative embodiment, the gate elements can be segmented, thus each segment of the grid elements can be operated independently, i.e. open and close at different timing, as a segmented ion gate. The method for manufacturing an ion gate for a charged particle stream, can comprise of electrically isolated grid elements that lie in the same plane. The respective sets of grid elements can be applied to alternate potentials to close the gate and the same potential to open the gate. For example, the grid element is at **100** volts above and **100** volts below the reference potential. The reference potential is the potential at the particular location of the gate in the drift tube. The steps used to manufacture the ion gate can be in any order and comprise: fabricating at least two gate elements, wherein each gate element includes at least one set of grid elements; assembling an insulating layer between the gate elements; deflecting at least one set of grid elements into the same plane of the other set of grid elements with at least one substrate; and securing the gate elements and the non-conductive substrates together.

A non-limiting example of the completed ion gate is shown in FIG. 1. The shape of the ion gate can be: square (shown), oval, circle, semicircle, triangle, rectangle, polygon, octagon, but not limited to these examples. FIG. 1 shows a front view of the completed ion gate. The ion gate includes gate elements that contain the grid elements **103** are held in place with several screws **105**. The ion gate apparatus that is used for gating a charged particle stream comprises: at least two sets of grid elements that individual voltages can be applied to each set to open and close; and at least one substrate for deflecting at least one set of grid elements into the same plane of the other sets of grid elements. In addition an electrically insulating layer can be added to allow different voltages to be set to each set of grid elements.

Another aspect of the invention relates to providing an ion gate with an effective gating function by applying a uniform tension on the grid elements, fabricating the gate elements such that the grid elements are equally spaced, and deflecting the grid elements into the same plane.

FIG. 2 shows a cross-sectional top view of the ion gate. The cross-section shown in FIG. 2 is indicated in FIG. 1 with a cutting line **107**. The ion gate comprises a first and second electrically isolated gate elements **204** and **206** that each have at least one grid element **205** and **207**. The electrically non-conductive or conductive substrate **208** deflects the grid element **207** which is part of the gate element **206** into the same plane as the deflected grid element **205** which is part of the gate element **204**. The gate elements **204** and **206** are electrically isolated by placing an insulating layer (electrically non-conductive or conductive) **210** between these gate elements. The grid elements are deflected into the same plane **202** and are electrically isolated by interleaving these sets of grid

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elements with a gap between each grid element. FIG. 5C illustrates the respective sets of grid elements in FIG. 5A and FIG. 5B interleaved. In FIG. 2, the ion gate components are secured together with a screw fastener 212. In this non-limiting example, either an electrically non-conductive material

screw fastener may be used or a conductive material fastener can be contained in an electrically non-conductive standoff (not shown). One embodiment of the present invention, involves using off-set electrically non-conductive or conductive substrates to deflect the grid elements. FIG. 3A shows the electrically

substrates 305 and 307 with no off-set 301. FIG. 3B shows electrically substrates 305 and 307 with an off-set 303. This off-set design may allow uniform deflection of grid elements and low cost manufacturing. Another embodiment of the present invention, involves the shape of the electrically non-conductive or conductive substrate to deflect the grid elements. The portion that deflects the grid element can be in the shape of a wedge, hexagon, semi-circle, but not limited to these examples. In addition, the electrically non-conductive substrate portion that deflects the grid element can be independent to the secured electrically non-conductive substrate. A non-limiting example is shown in FIG. 4, where a circular (not limited to only this shape) substrate 402 deflects the grid element and the secured substrate 404 holds the substrate 402 in place.

Yet another embodiment of the present invention is the fabrication of the gate elements. The grid elements within the gate elements can be produced by cutting, etching, evaporation or electroplating, but not limited to these methods. In a non-limiting example, parallel rows of grid elements are formed by removing portions of a given thickness of planar metal foil by etching portions from the foil. This method forms a plurality of grid elements that are equally spaced. In addition to forming equally spaced grid elements within the gate element, the grid elements are made from the same gate element material as a single entity. In this manner, the grid elements do not need to be fixed to the gate element through gluing (epoxy), glass soldering, or any other attaching manner. Fabricating the gate elements by etching the grid elements is a robust and reproducible method for manufacturing the grid elements. In addition, since no gluing, soldering, or other attaching manner is used in fabricating the gate elements, elevated temperatures and/or thermal expansion of the grid elements are all uniform. The photo chemical milling (etching) can be performed on one side or both sides of the material being etched.

FIG. 7A shows the shape (a pair of opposing concave fillets) of the grid element etched on both sides of the material, etched from the top 701 and the bottom 702 of the material. FIG. 7B shows the shape (a concave fillet) of the grid element etched on one side of the material, etched from only the top side 704 of the material. The etched edge distance 703 may be smaller or greater than the material thickness 705 depending on the layout of the grid elements. For example, the etched edge distance to the material thickness ratio is greater than zero, in particular 1-50%, 50-100%, 100-500%. FIG. 7C shows three grid elements that are etched on both sides of the material. Photo chemical milling can produce uniform dimensions in grid element width 707 and gap 709 between grid elements forming a plurality of grid elements that are equally spaced. When the ion gate is a uniform product, injection of ion packets into the drift tube are tight packets with limited background signal, therefore a higher signal to noise ratio can be achieved. In this embodiment, regardless of manufacturing methods, the grid element is to be made with a sharp edge, the geometry may generate a narrow gating

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electric field region resulting in high precision gating of charge particles. In an IMS device, a narrow ion pulse could be generated with precision gate timing control. An ion gate apparatus for gating a charged particle stream comprises: at least two sets of electrically insulated grid elements on the same plane, the evenly spaced grid elements have at least one sharp edge face to adjacent grid element.

Another embodiment of the present invention is securing the gate elements together with non-conductive substrates and the insulating layer. The ion gate can be secured by clamping, soldering, screws, pins, but not limited to these examples. The insulating layer can be made from any non-conductive material such as, ceramic, aluminum nitrate, but not limited to these examples.

An alternative embodiment of manufacturing an "Tyndall" type of ion gate for a charged particle stream involves two sets of electrically isolated grid elements, wherein the first set of grid elements is arranged with an offset in respect to the second set of the grid elements, such that the gaps of the first grid element is aligned with the second grid elements; each set of grid elements is applied to alternate potentials when the gate is closed and same potential when the gate is opened. The method involves the steps of: fabricating at least two gate elements by removing portions of a substantially planar metal foil to form a plurality of grid elements; wherein each gate element includes at least one set of grid elements; assembling an insulating layer between the gate elements; and securing the gate elements together.

In another embodiment of the invention, a metalized dielectric structure is used for the Bradbury-Nielsen gate. A ceramic material is coated with single or multiple layers of metallization materials. The metallization process is commonly finished with a thin layer of nickel, gold or other inert metal for enhanced chemical resistivity. FIG. 8 shows a unique construction method of a Bradbury-Nielsen ion gate using a metalized ceramic material. It is built with a frame ring, a tension ring 802 and parallel wires 805 that are pre-winded on a metal frame. One or the rings, either the frame ring or the tension ring is metalized 815 with a pattern 810 that connects every other wire to each other. In one embodiment, the frame ring has metalized contacts 807 that are 1 mm apart (center to center). During the ion gate construction, the parallel wires are lined up with these contacts and form a firm contact while the tension ring is pushed down into the frame ring. As the wire is selected to match the thermal expansion of the frame ring and tension ring, the wires can be maintained parallel while the IMS is operated under different temperature conditions. The gate control voltage(s) are applied to the wires by attaching an electrical lead to the contact point that is on the outside of the frame ring. Not only for metalized ceramic tube IMS design, the Bradbury-Nielsen ion gate can be used for other analytical instruments.

In various embodiments for symmetric IMS, an ion focusing method can be employed to guide ions to a target collection area on the collector. Suitable focusing methods may include, but are not limited to, static electric field focusing and ion funnel focusing. An ion collector can be segmented to facilitate, collection of ions with specific ion mobility (drift time) or a certain range of mobilities on to different segment of the ion collectors. A segmented Bradbury-Nielsen gate can be used to enhance the separation and collection.

In various embodiments of IMS instruments, wherein the Bradbury-Nielsen gate can be segmented. A variety of geometries, including but not limited to parallel, rectangular, concentric ring shape, can be used for the segmentation, referring to FIG. 9, various embodiments can use parallel segmentation. Each segment of the ion gate, for example, 901, 903, and

905, can be controlled to open at a different time. Such a segmented ion gate can be used as either first or second ion gate in a time-of-flight type ion mobility separator. While it is used as the second ion gate in a IMS, multiple portions of ions with different drift time are allow to pass through segmented ion gate, thus collected on different sections of ion collectors, and recovered separately if desired.

In various embodiments, an apparatus of ion gate for an ion mobility separator comprising a segmented Bradbury-Nielson gate that contains multiple sections of Bradbury-Nielson gate. The segmented Bradbury-Nielson gate can be used as a second gate in a time-of-flight type ion mobility separator. The segmented Bradbury-Nielson gate comprises a variety of geometries which may include but is not limited to: parallel, rectangular, concentric. The ion mobility separator further comprises a segmented ion collector where a plurality of sections of ion collector is inline with the sections of the segmented Bradbury-Nielson gate.

This invention further describes a method and apparatus of ion gate operation. In one embodiment, an AC voltage is used to close the gate. In a common operation of the Bradbury-Nielson gate, the ion gate is open when the adjacent grid elements are at the same potential and the ion gate is closed when a DC voltage, e.g. 30V, 50V, 100V, 200V, or -30V, -50V, -100V, -200V are applied on the adjacent grid elements. The voltage creates an electric field that pushes ions toward the grid element that is a lower potential, thus preventing ions from penetrating through the ion gate when closed.

During ion mobility measurements, the ion gate is opened for a short period of time, e.g. 100 microseconds, and then closed for a period of time, e.g. 20 millisecond, while ions are traveling in the drift tube. FIG. 10 illustrates a closed ion gate. When the applied voltage is higher than the reference potential, a '+' is shown for the gate wire (i.e. grid element) and when the applied voltage is lower than the reference potential, a '-' is shown for the gate wire. The voltage difference between the adjacent wires causes positive ions to be collected on the (-) wire and negative ions to be collected on the (+) wire. By repeating wire layout pattern of (+) and (-) wires, a large area is covered and ions are prevented to pass through the ion gate. In this embodiment, an AC voltage instead of DC voltage is applied the gate wires. In this case, the potential of each wire is constantly changing. The potential on wire 1003 is '+' and wire 1004 is '-' at the state 1001, and then the potential on wire 1003 is '-' and wire 1004 is '+' at the state 1002. The state 1001 and 1002 repeats at certain frequency. The voltage and frequency of the AC applied to the adjacent wires is optimized to completely close the ion gate. The ion gate is opened by setting all gate wires at the same potential. The new gate configuration and operational method is not limited to be used for ion mobility spectrometer, but can be used for any device that needs to shut off a stream of charged particles.

In a variety of the embodiments, a method for operating an ion gate for a charged particle stream involves opening the ion gate by setting adjacent gate wires (grid elements) at the same potential; closing the ion gate by setting adjacent gate wires at different potentials by applying an AC voltage to the adjacent wires. The AC voltage can be controlled to provide a given frequency and amplitude that that is most suitable for the intended operation. The frequency can be a constant and/or controlled to cover a broad range during a period when the ion gate is closed. Similarly, the amplitude can be a constant and/or controlled in a range during a period of closing the gate. The AC voltage may be a symmetric or asymmetric waveform (for example, a waveform is used for DMS and/or FAIMS, a typical example of the waveform is shown in FIG.

21). In a variety of operation modes, the AC powered ion gate can be operated as such, conducting a series of ion mobility measurements under a series of different frequencies and/or amplitudes, and then integrating the series of ion mobility measurement data into an ion mobility spectrum using common data processing algorithms, such as summing.

A Bradbury-Nielson gate is traditionally used for injecting short pulses of ions into TOF mass spectrometers and ion mobility spectrometers to improve the mass resolution of TOF instruments by reducing the initial pulse size as compared to other methods of ion injection. Therefore the precise control of the ion pulse width admitted to the drift tube can be controlled. However, the means for controlling the size of ions entering the drift tube using a Bradbury-Nielson gate has not been proposed to date.

One embodiment of the present invention is to use an AC voltage instead of the commonly used DC voltage to control the amount of a ion pulses generated and let through the gate. In a common operation of the Bradbury-Nielson gate, the ion gate is open when the adjacent grid elements are at the same potential and the ion gate is closed when a DC voltage, e.g. 30V, 50V, 100V, 200V, or -30V, -50V, -100V, -200V are applied on the adjacent grid elements. The voltage creates an electric field that pushes ions toward the grid element that is a lower potential, thus preventing ions from penetrating through the ion gate when closed. FIG. 11 shows an example of a gate using DC voltage. Negative sample ions 1101 travel toward the ion gate wires 1105 and 1106 of the gate in a relatively straight line according the applied field in the reaction region (desolvation region) 1103 until they come close to the gate wires where they are pulled toward the positive wires 1105 and are neutralized on the positive wires 1105 if the gate is not in the open state. Since DC voltage is being used to control the gate, each adjacent wire has a different polarity, either positive or negative. Therefore gate wires 1105 are positive and gate wires 1106 are negative. During ion mobility measurements the ion gate is opened for a short period of time, e.g. 100 microseconds, and then closed for a period of time, e.g. 20 millisecond, thereby letting a pulse of ions travel in the drift tube 1108. During the closed state of the gate an ion depletion area is formed 1110. The larger the area of the ion depletion area the longer the gate needs to be opened to let ions through, since there is a low population of ions. In order to get higher resolution peaks in the IMS spectrum, a shorter opening period of time is preferred in order to get a narrow pulse of ions through with limited diffusion. By using an AC voltage instead of a DC voltage to close the gate the ion depletion area can be reduced, in particular, by a half. The ion depletion area can be reduced by an percentage by using an AC voltage.

In a variety of the embodiments, a method for operating an ion gate for a charged particle stream involves opening the ion gate by setting adjacent gate wires (grid elements) at the same potential; closing the ion gate by setting adjacent gate wires at different potentials by applying an AC voltage to the adjacent wires. The AC voltage can be controlled to provide a given frequency and amplitude that that is most suitable for the intended operation. The frequency can be a constant and/or controlled to cover a broad range during a period when the ion gate is partially open and/or closed. Similarly, the amplitude can be a constant and/or controlled in a range during a period of closing the gate. The AC voltage may be a symmetric or asymmetric waveform. The waveform can be, but not limited to: sine, square, triangle, and sawtooth. In a variety of operation modes, the AC powered ion gate can be operated as such, conducting a series of ion mobility measurements under a series of different frequencies and/or amplitudes, and then

integrating the series of ion mobility measurement data into an ion mobility spectrum using common data processing algorithms, such as summing. FIG. 12 shows an example of a gate using AC voltage. Negative sample ions **1201** travel toward the ion gate wires **1205** of the gate according to the AC applied field until they come close to the gate wires where they are pulled toward the gate wires **1205** and are neutralized if the gate is not in the open state. Since AC voltage is being used to control the gate, each adjacent wire has a different polarity, which alternates between positive or negative. Therefore the gate wires **1205** are positive and are negative according to the applied AC voltage. During ion mobility measurements, the ion gate is opened for a short period of time, e.g. 100 microseconds, and then closed for a period of time, e.g. 20 millisecond, while ions are traveling in the drift tube **1208**. During the closed state of the gate an ion depletion area is formed **1210**. By using an AC voltage instead of a DC voltage to close the gate the ion depletion area is minimized, in particular, can be reduced by a half thereby improving the operation of the ion gate. There is little or no space charge using the AC voltage as compared to DC voltage. Since the ion gate can create tighter ion pulses when it is opened the resolution of the peaks traveling through the drift tube **1208** are significantly increased. FIG. 13 shows the AC gate in the closed position **1301** and then completely open **1303** and then closed again **1305**. The open **1303** position is for a shorter period of time than that when using a DC voltage.

Another embodiment of the present invention is to use an AC voltage instead of DC voltage on an ion gate to filter ions according to their size and/or ion mobility. An apparatus and method is used to filter ions according to their ion mobility by varying the AC potential on a segmented Bradbury-Nielson gate. Some of the components of the charged particle stream have a smaller size and/or shape (or ion mobility) compared to the other components of the charged particle stream. The present invention controls the AC voltage on a segmented Bradbury-Nielson gate in order to filter small ions from larger ions from an ionized sample before they enter the drift tube. FIG. 14 shows an example of an IMS spectrum using 20 parts per million (ppm) of an L-Tryptophan sample in 80/20 methanol and water using a DC voltage on the ion gate. The reactant ion peak (RIP) and L-Tryptophan (L-Trp) are shown as peaks in the spectrum. When AC voltage is used on the ion gate smaller sized ions such as the RIP peaks can be substantially eliminated as shown in FIG. 15. The AC frequency can be tuned to control the specific ion mobility that is excluded from being transmitted from the ionized sample into the drift tube. For example, an AC frequency of 67 kHz and a 100-240 voltage range between the gate wires would exclude small ions having a 100 molecular weight size from the other ions in the sample that have a larger molecular weight greater than 100. FIG. 16 shows the AC mobility selecting gate in the closed position **1601** having a voltage of 240 and then partially open **1603** having a voltage of 100 and then closed again **1605** having a voltage of 240. In this mode, when the gate is open **1603**, only larger molecular weight (low ion mobility) components of the sample can get through the gate without getting neutralized on the gate wires. The AC mobility selecting gate works by tuning the AC frequency in the reaction region (desolvation region) **1705** as shown in FIG. 17. When components of a sample have different ion mobilities, sizes or molecular weights, the AC frequency has a stronger or weaker effect on the components travel. For example, for two negative ions where one **701** has a molecular weight of 100 and the other ion **1703** has a molecular weight of 200 the trajectory using a 67 kHz frequency is substantially different as shown in FIG. 17. The smaller (high mobility) ion **1701** has a larger

oscillation **1708** than the larger (low mobility) ion's oscillation **1706**. When the AC gate is only partially open as in FIG. 16 the larger ion **1703** passes the gate wires onto the drift tube **1708** and the smaller ion **1701** is neutralized on the gate wire.

In yet another embodiment, the wires of the AC gate can be configured to enhance the size selection. As shown in FIG. 18, the wires **1815** can be made rectangular such that certain small ions would have a larger surface area in the direction that is parallel to the charged particle stream to be neutralized on and the frequency tuning could be established such that the small and large ions are within a few molecular weight units and still be differentiated. Alternatively, the wires **1815** in FIG. 18 can be 2 or more that are aligned in phase as shown in FIG. 19 or 2 or more are out of phase in any array as shown in FIG. 20. When 2 or more wires are used in an array that are out of phase a specific component ion mobility can be filtered out thereby letting the other components through the gate. The configuration shown in FIG. 20 provides the ability eliminate one specific sample component from traveling through the gate to the drift tube **2008**. This is different from other configurations where they operate as a cut-off whereby all small size and/or shape ions are eliminated at a certain point. In a variety of the embodiments, a method for operating an AC ion gate for charged particles stream involves, applying an AC voltage to 2 or more wires; the AC voltage are out of phase, selectively eliminating one or more sample components from the charged particle stream while partially opening the ion gate; conducting ion mobility and/or mass measurement. The operating method may further include applying a series of AC voltages in certain sequence during ion mobility measurement. Such sequence may include AC voltage that selectively eliminating sample component ions in the charge particle stream from high ion mobility to low ion mobility; or from low mobility to high mobility; or selectively for one or more targeted sample components with certain ion mobilities. The frequency and/or amplitude of the AC voltages in such sequence may be altered in a continuous or discrete fashion.

In a variety of the embodiments, a method for operating an AC ion gate for a charged particle stream involves; applying an AC voltage to one or more of the gate wires to partially open and/or close the ion gate, partially opening the ion gate with an AC voltage that is greater than 0 but less than the AC voltage used to close the ion gate, and filtering a percentage of some components of the charged particle stream from the other components of the particle stream by neutralizing such components on the ion gate wires. The frequency and/or amplitude of the AC voltage can be controlled in a range or held constant during a period of closing the gate. The AC voltage can be a symmetric or asymmetric waveform. The percentage of some components of the charged particle stream being filtered from the other components can be 0 to 100%, in particular, 100%, greater than 75%, greater than 50%, greater than 25%, greater than 10%, or a small percentage being greater than 0%.

In one embodiment, the method of operating an AC gate may include a series of pulse of ions that allow different components of charged particle stream to pass through the AC gate; and analyze the components of charged particle stream based on their ion mobility and/or mass to charge ratio.

In one preferred embodiment of a time of flight ion mobility spectrometer include an ionization source located on one end of the ion mobility spectrometer is used to ionize the samples. An electric field is used to guide the ionized sample toward an ion gate. On the ion gate, at least one AC voltage is applied to the gate elements where at least one pulse of the ionized sample is passed into a drift tube. The ionized sample is then guided by another electric field to guide the pulse(s) of

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ions toward an ion detector. The ion detector is commonly located at the other end of the spectrometer. The AC voltage can be composed using one or more waveform. The AC voltage has a waveform that is substantially same as the asymmetric waveform used in differential/field asymmetric ion mobility spectrometers, an example of such waveform is shown in FIG. 21. The time of flight ion mobility spectrometer can offer two separation mechanisms in one integrated structure. The ionized sample mixture is first filtered by the AC ion gate based on their ion mobility differential under high electric field condition between grid elements; only selected ions under a given AC voltage condition can pass the ion gate as a pulse of ions, and then, the pulse of ions are further separated in the drift tube of the time of flight ion mobility spectrometer based on their low field ion mobility. The AC gate can sweep through a range of AC voltages to select ions with different high field mobility to be pulsed into the drift tube for further separation. With this measurement, a 3D separation plot could be generated with one axis of low field ion mobility, or drift time, and another axis of compensation voltage of the asymmetric waveform for DMS/FAIMS, and the third as axis of ion intensity.

Even though many embodiments and examples given in this disclosure refer to ion gate for general IMS device, these devices can be operated under low vacuum, ambient or high pressure conditions. Alternatively, the ion gate can be operated in liquid for liquid phase IMS or other devices, such as electrophoretic devices, where packets of ions need to be formed. The ion gate can also be used under vacuum conditions for generating ion packets for mass spectrometers, such as a time of flight mass spectrometer. This invention discloses gating methods and apparatuses that can be used for any device where packets of charged particles need to be formed.

It is recognized that modifications and variations of the invention disclosed herein will be apparent to those of ordinary skill in the art and it is intended that all such modifications and variations be included with the scope of the appended claims.

What is claimed is:

1. An apparatus of an ion gate for a charged particle stream in a spectrometer comprising:
 - (a) at least one pair of adjacent grid elements that can be set at the same or different potentials;
 - (b) at least one AC voltage is applied to the grid elements to pass a pulse of selected ions into the spectrometer; wherein the pulse of selected ions is generated by applying a first frequency and a first amplitude of the AC voltage to at least one of the grid elements for a period time and subsequently applying a second frequency and/or a second amplitude of the AC voltage to at least one of the grid elements.
2. The apparatus of claim 1, wherein the grid elements have a large surface area in the direction that is parallel to the charged particle stream.
3. The apparatus of claim 1, wherein the grid elements are 2 or more that are aligned in phase.

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4. The apparatus of claim 1, wherein the grid elements are 2 or more that are aligned out of phase.

5. A method for operating an AC ion gate for a charged particle stream comprising: Applying at least one AC voltage to at least one of the grid elements of the AC ion gate to pass a pulse of selected ions into the spectrometer; wherein the pulse of selected ions is generated by applying a first frequency and a first amplitude of the AC voltage to at least one of the grid elements for a period time and subsequently applying a second frequency and/or a second amplitude of the AC voltage to at least one of the grid elements.

6. The method of claim 5, wherein the pulse of selected ions is generated by partially opening the ion gate using a selected AC voltage.

7. The method of claim 6, further comprising: filtering a percentage of some components of the charged particle stream from the other components of the particle stream by neutralizing such components on the grid elements.

8. The method of claim 7, wherein the percentage of some components of the charged particle stream being filtered from the other components is 0 to 100%: in particular, 100, >75, >50, >25, >10, and >0.

9. The method of claim 7, wherein some components of the charged particle stream have a smaller size and/or shape than the other components of the charged particle stream.

10. The method of claim 7, wherein the other components of the charged particle stream have a larger size and/or shape than some components of the charged particle stream.

11. The method of claim 5, wherein the amplitude can be a constant and/or controlled in a range during a period of closing the gate.

12. The method of claim 5, wherein the AC voltage is a symmetric or asymmetric waveform.

13. The method of claim 5, wherein the frequency can be a constant and/or controlled in a range during a period of closing the gate.

14. A time of flight ion mobility spectrometer comprising:

- (a) an ionization source that ionize samples
- (b) a first electric field that guides the ionized sample toward an ion gate
- (c) at least one AC voltage is applied to the grid elements of the ion gate for a period of time, by which at least one pulse of the ionized sample is passed into a drift tube
- (d) a second electric field that guides the pulse(s) of ions toward an ion detector.

15. The time of flight ion mobility spectrometer of claim 14, wherein the AC voltage has a waveform that is same as the asymmetric waveform used in differential/field asymmetric ion mobility spectrometers.

16. The time of flight ion mobility spectrometer of claim 14, wherein the AC voltage is a controlled sweep through a range of voltages and/or frequencies to select different sample ions to pulse into the drift tube.

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